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OCT 10 2006 Amendment

Application No.10/511,527

AMENDMENTS TO THE CLAIMS

This listing of claims replaces all prior versions, and listings, of claims in the application.

1. (Previously Presented) Method for the detection and characterisation of primary tumours and separate areas of primary tumours, respectively, method comprising using sample material to isolate and concentrate cell clusters of tumour cells, followed by an analysis of the genetic changes in these isolated cell clusters.
2. (Previously Presented) Method according to claim 1, sample material consists of cell cultures, blood, urine, nipple aspiration fluid from the female breast or tissue from primary tumours.
3. (Previously Presented) Method according to claim 1, wherein polymorphic DNA of primary tumours or separate areas of primary tumours, and alterations therein, respectively, are recorded and compared with corresponding polymorphic DNA of cell clusters, and alterations therein, respectively.
4. (Previously Presented) Method according to claim 1, wherein DNA of the following polymorphic sequences are analysed: D7S522, D8S133, D8S258, D8S265, NEFL, D10S541, D10S1765, D10S579, D13S153, D16S400, D16S402, D16S413, D16S422, p53, BB1, BB2, CAlI, CAII, CAIV, CAV and/or D17S855.
5. (Previously Presented) Method according to claim 1, wherein the polymorphic DNA is reproduced before analysis.
6. (Currently Amended) Method according to claim 5, wherein the polymorphic DNA of three polymorphic sequences, D7S522, ~~D8S256~~ D8S258, D16S400 or NEFL, D13S153, D17S855 or D10S541, D16S402, D16S422 are analysed together and/or reproduced.
7. (Previously Presented) Method according to claim 6, wherein the polymorphic DNA is reproduced prior to analysis by polymerase chain reaction (PCR).

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8. (Previously Presented) Method according to claim 7, wherein the polymorphic DNA is reproduced by using the following primer pairs:

GCAGGACATGAGATGACTGA and GTTATGCCACTCCCTCACAC (for D7S522);
GTTTGAAGAATTTGAGCCAACC and TTCTTCTGCACACTTGGCAC (for BB1+2);
CTCGAGGTCTCATCCTCTTTCC and GCAGAGGTGCACAAAGGAGTAA (for CAII);
AGGCCACAGAGGAGATAACAG and CAGGTGTGGTAGATGCCAAAGA (for CAIII);
GCAACTTATCCAAACCCTGACC and AGAGTGGACTAGGAAATGCTAGGAG (for CAIV);
AGTTCCTGACTGGGAATTCGAT and TTGGCCAAATTACACACCTTTG (for CAV);
TTCCATTTGTCICGGTT and AGTCTCCTCGTCTCACACCT (for D7S2550);
CAGTGCTGGAGTTGTTCAAG and CTGGGAGTCAAGTGTTTGG (for D7S2429);
TGCTAAGTCTTGATTTTGCC and AACGGTCATCTGTGTTGG (for D7S2467);
GGTGTTTGTGTCAATTACGCT and TTTGCTGTAGAGGATGCAAT (for D7S478);
TTCGGCCTCTCTGTTATAAA and CCGAAGCAGGATTTTATTC (for D7S670);
AGCTGCCAGGAATCAACTGAGAG and GATGCTCACATAAAGGAGGGAGG (for D8S258);
CCAATACCTGCAGTAGTGCC and GAGCTGCTTAACACATAGGG (for NEFL);
CACCACAGACATCTCACAACC and CCAGTGAATAGTTCAGGGATGG (for D10S541);
AGGGTTATGTATAACCGACTCC and GTCTAAGCCCTCGAGTTGTGG (for D13S153);
GTTTCACAAATTGGACAGTAT and GAACCCTCCATGCTGACATT (for D16S400);
GTACCCATGTACCCCCAATA and CAAAGCACCATAGACTAA (for D16S402);
GAGAGGAAGGTGGAAATACA and GTTTAGCAGAATGAGAATAT (for D16S422);
AATAAATTCCCACTGCCACTC and ATCCCTGAGGGATACTATTC (for p53);
GGATGGCCTTTTAGAAAGTGG and ACACAGACTTGTCTACTGCC (for D17S855).

9. (Previously Presented) Method according to claim 5, wherein the reproduced DNA fragments are split and analysed by capillary electrophoresis.

10. (Previously Presented) Method according to claim 1, wherein the isolation or concentration of tumour cells cytokeratin-positive cells were isolated from sample material, and/or positive epithelial cells for tissue specific proteins.

11. (Previously Presented) Method according to claim 10, wherein epithelial cells are concentrated from sample material by means of density gradient centrifugation-if

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necessary after homogenisation in a solvent, and cytokeratin-positive and/or positive cell clusters from tissue specific proteins are then split off by means of immunomagnetic cell isolation.

12. (Previously Presented) Method according to claim 11, wherein the medium for the density gradient centrifugation is a hyper-osmotic medium.

13. (Previously Presented) Method according to claim 12, wherein the hyper-osmotic buffer consists of one of the following mediums: 13.8% (w/v) Diatrizoate and 8% (w/v) dextran 500 in H₂O (polymorphprep) or 13% (w/v) Nycodenz, 0.58% (w/v) NaCl and 5 mM Tricine-NaOH pH 7.4 in H₂O (Nycoprep).

14. (Previously Presented) Method according to claim 1, wherein genetic changes in the isolated cell clusters are analysed by means of cluster analysis.

15. (Previously Presented) Application of a method according to claim 1 for the molecular characterization of tumours or tumour sections or for the determination of clonality from cells clusters isolated from sample material as well as for the detection of a tumour to determine the tumour stage, the metastasising potential, therapy requirements, efficacy of therapy of a tumour or part thereof, as well as the assessment of the course of a disease or therapy.

16. (Previously Presented) Application according to claim 15 for the detection and/or characterisation of tumours or tumour areas of the following carcinomas: mamma-, ovarian-, colon-, gastric-, prostate and/or bladder carcinoma.